

PATENT COOPERATION TREATY

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INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

REC'D 03 JAN 2005

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Applicant's or agent's file reference 2004647-0009	FOR FURTHER ACTION See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416)	
International application No. PCT/US03/06491	International filing date (day/month/year) 05 March 2003 (05.03.2003)	Priority date (day/month/year) 06 March 2002 (06.03.2002)
International Patent Classification (IPC) or national classification and IPC IPC(7): G01N 33/567 and US Cl.: 435/7.2		
Applicant TRUSTEES OF BOSTON UNIVERSITY		

1. This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36.

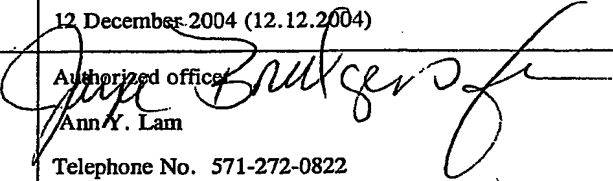
2. This REPORT consists of a total of 3 sheets, including this cover sheet.

☐ This report is also accompanied by ANNEXES, i.e., sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT).

These annexes consist of a total of ___ sheets.

3. This report contains indications relating to the following items:

- I ☒ Basis of the report
- II ☐ Priority
- III ☐ Non-establishment of report with regard to novelty, inventive step and industrial applicability
- IV ☐ Lack of unity of invention
- V ☒ Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- VI ☐ Certain documents cited
- VII ☐ Certain defects in the international application
- VIII ☐ Certain observations on the international application

Date of submission of the demand 06 March 2003 (06.03.2003)	Date of completion of this report 12 December 2004 (12.12.2004)
Name and mailing address of the IPEA/US Mail Stop PCT, Attn: IPEA/US Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450 Facsimile No. (703) 305-3230	Authorized officer  Ann Y. Lam Telephone No. 571-272-0822

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

PCT/US03/06491

I. Basis of the report

1. With regard to the elements of the international application:*

- ☒ the international application as originally filed.
- ☒ the description:
pages 1-104 _____ as originally filed
pages NONE _____, filed with the demand
pages NONE _____, filed with the letter of _____.
- ☒ the claims:
pages 105-120 _____, as originally filed
pages NONE _____, as amended (together with any statement) under Article 19
pages NONE _____, filed with the demand
pages NONE _____, filed with the letter of _____.
- ☒ the drawings:
pages 1-9 _____, as originally filed
pages NONE _____, filed with the demand
pages NONE _____, filed with the letter of _____.
- ☐ the sequence listing part of the description:
pages NONE _____, as originally filed
pages NONE _____, filed with the demand
pages NONE _____, filed with the letter of _____.

2. With regard to the language, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item.

These elements were available or furnished to this Authority in the following language _____ which is:

- ☐ the language of a translation furnished for the purposes of international search (under Rule 23.1(b)).
- ☐ the language of publication of the international application (under Rule 48.3(b)).
- ☐ the language of the translation furnished for the purposes of international preliminary examination (under Rules 55.2 and/or 55.3).

3. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:

- ☐ contained in the international application in printed form.
- ☐ filed together with the international application in computer readable form.
- ☐ furnished subsequently to this Authority in written form.
- ☐ furnished subsequently to this Authority in computer readable form.
- ☐ The statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.
- ☐ The statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished.

4. ☐ The amendments have resulted in the cancellation of:

- ☐ the description, pages NONE
- ☐ the claims, Nos. NONE
- ☐ the drawings, sheets/fig NONE

5. ☐ This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed, as indicated in the Supplemental Box (Rule 70.2(c)).**

* Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17).

** Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this report.

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.
PCT/US03/06491**V. Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement****1. STATEMENT**

Novelty (N)	Claims <u>NONE</u>	YES
	Claims <u>1-85</u>	NO
Inventive Step (IS)	Claims <u>NONE</u>	YES
	Claims <u>1-85</u>	NO
Industrial Applicability (IA)	Claims <u>1-85</u>	YES
	Claims <u>NONE</u>	NO

2. CITATIONS AND EXPLANATIONS

Claims 1-85 lack novelty under PCT Article 33(2) as being anticipated by Thalhammer-Reyero, US 5,930,154A. Thalhammer-Reyero discloses a method of constructing a model of a biological network comprising: providing a biological system comprising a plurality of biochemical species, including genes, polypeptides and metabolites, (col. 7, lines 30-40); perturbing the activity of the biochemical species, causing a response in the biological network (col. 15, lines 7-18); allowing the biological network to reach a steady state (col. 7, lines 30-33); determining the response of at least one of the biochemical species in the biological network (col. 15, lines 7-18); and estimating parameters of the model (col. 5, lines 1-7.) Equations that represent evolution over time of the activities is disclosed (col. 16, lines 45-49, col. 14, lines 7-18, and col. 41, lines 44-56.) Measuring the activity of the biochemical species is disclosed (col. 15, lines 52-58). Alterations in the activity of the biochemical species is disclosed (col. 41, lines 53-56.) A fitness function to predict the measured values of the perturbation is disclosed (col. 41, lines 49-56.) Imposing constraint on the biological network is disclosed (col. 41, lines 49-61.) Calculating or searching for parameters that optimize a chosen fitness function, and stop criterion, are disclosed (col. 41, lines 54-65.) A density function for each estimated parameter is disclosed (col. 41, lines 43-46.) Determining the sensitivity of the activity of one species to a change in the activities of another species, wherein the measure is a quantitative measure is disclosed (col. 7, lines 30-34, col. 41, lines 44-56.) A method of identifying a target of a perturbation, and identifying phenotypic mediators is disclosed (col. 7, lines 30-34, and 41-45.) A computer system for constructing a model of a biological network, including a memory and a processor using a code system, wherein the computer calculates parameters and identifies a target of a perturbation or a phenotypic mediator is disclosed (col. 15, lines 27-64, col. 31, lines 15-26, col. 32, lines 21-29, col. 42, lines 8-15.)